## CLAIMS

We claim:

- 1. An immunoglobulin comprising a protection protein in association with an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain.
- 2. The immunoglobulin of claim 1 further comprising an immunoglobulin derived light chain having at least a portion of an antigen binding domain associated with said immunoglobulin derived heavy chain.
- 3. The immunoglobulin of claim 1 or 2 further comprising a second immunoglobulin derived heavy chain 15 having at least a portion of an antigen binding domain associated with said protection protein.
- 4. The immunoglobulin of claim 3 further comprising a second immunoglobulin derived light chain having at
  least a portion of an antigen binding domain bound to said
  20 second immunoglobulin derived heavy chain.
  - 5. The immunoglobulin of claims 1-4 further comprising immunoglobulin chain bound to at least one of said immunoglobulin derived heavy chains.
- 6. The immunoglobulin of claims 1-5 that is a 25 therapeutic immunoglobulin.
  - 7. The immunoglobulin of claim 6 wherein said therapeutic immunoglobulin binds to mucosal pathogen antigens.
- 8. The immunoglobulin of claim 7 that is capable 30 of preventing dental caries.
  - 9. The immunoglobulin of claims 1-8 wherein said antigen binding domain is capable of binding an antigen from <u>S. mutans</u> serotypes c, e and f or *S. sobrinus* serotypes d and g.
  - 10. The immunoglobulin of claim 1 wherein said protection protein has an amino acid sequence which substantially corresponds to at least a portion of the

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amino acid residues 1/to 627 of the rabbit polyimmunoglobulin receptor and does not have an amino acid residue sequence corresponding to amino acid residues 628-755 of the rabbit polyimmunoglobulin receptor.

- The immunoglobulin of claim 1/ wherein said protection protein has an amino acid /sequence which substantially corresponds to at least a portion of the amino acid residues 1 to 606 of the rabbit polyimmunoglobulin receptor and does not have an amino acid sequence corresponding to amino acid residues 628-755 of the rabbit polyimmunoglobulin receptor.
- The immunoglobulin of glaim 10 or 11 wherein said protection protein has an amino acid sequence which does not contain amino acid residues corresponding to amino acid residues 628, ťο 775 of the polyimmunoglobulin receptor/and which does contain amino acid residues which correspond to one or more of the following amino acid segments:
- amino acids corresponding to amino acid resia)
  - dues 21-43 of the rabbit polyimmunoglobulin receptor;
    b) amino acids corresponding to amino acid residues 1 - 118 of the rabbit polyimmunoglobulin receptor;
  - amino acids corresponding to amino acid residues 119 - 223 of the rabbit polyimmunoglobulin receptor;
  - amino acids corresponding to amino acid residues 224 - 332 of the rabbit polyimmunoglobulin receptor;
  - amino acids corresponding to amino acid residues 333 - 441 of the rabbit polyimmunoglobulin receptor;
  - amino acids corresponding to amino acid residues 442 - 552/of the rabbit polyimmunoglobulin receptor;
  - amino acids corresponding to amino acid resig) dues 553 606 or 553 - 627 of the rabbit polyimmunoglobu/in receptor.
- 13. The immunoglobulin of claim 1 wherein said protection protein has an amino dcid sequence which does 35 not contain amino acid residues of a polyimmunoglobulin receptor/of a species which are analogous to amino acid

residues 628 to 755 of the rabbit polyimmunoglobulin receptor and which does contain amino acid residues from a polyimmunoglobulin receptor of a species which are analogous to one or more of the following amino acid segments:

- a) amino acids corresponding to amino acid residues 21 43 of the rabbit polyimmunoglobylin receptor;
- b) amino acids corresponding to amino acid residues 1 118 of the rabbit polyimmunoglobulin receptor;
- c) amino acids corresponding to amino acid residues 119 223 of the rabbit polyimmunoglobulin receptor;
- d) amino acids corresponding to amino acid residues 224 332 of the rabbit polyimmunoglobulin receptor;
- e) amino acids corresponding to amino acid residues 333 441 of the rabbit polyimmunoglobulin receptor;
- f) amino acids corresponding to amino acid resi
  - dues 442 552 of the rabbit polyimmunoglobulin receptor;
    g) amino acids corresponding to amino acid resi-
- dues 553 606 or 553 627 of the rabbit polyimmunoglob-
  - 14. The immunoglobulin of claim 13 wherein said species is human.
  - protection protein includes the amino acid sequence of at least one of the domains selected from the group consisting of the following portions of the rabbit polyimmunoglobulin receptor: domain I, domain II, domain III, domain IV, domain V, and amino acid residues 553 to 627 of domain VI; and does not have an amino acid sequence corresponding to amino acid residues 628-755 of the rabbit polyimmunoglobulin receptor.
- 16. The immunoglobulin of claim 1 wherein said protection protein does not have any amino acid sequence which corresponds to or is analogous to amino acid residues 628-755 of the rabbit polyimmunoglobulin receptor and which does include:

- a) at least one domain which is from the polyimmunoglobulin receptor of a first animal and which is analogous to at least a portion of the following amino acid segments of the rabbit polyimmunoglobulin receptor: domain I, domain II, domain IV, domain V, and amino acid residues 553 to 627 of domain VI;
- b) at least one domain which is from the polyimmunoglobulin receptor of a second animal and which corresponds to or is analogous to the following amino acid
  residue segments of the rabbit polyimmunoglobulin receptor: domain I, domain II, domain III, domain IV, domain
  V, and amino acid residues 553 to 627 of domain VI.
- 17. The immunoglobulin of claim 1 wherein said protection protein does not have any amino acid sequence which corresponds to or is analogous to amino acid residues 628-755 of the rabbit polyimmunoglobulin receptor and which does include:
- a) at least one amino acid segment which is from the polyimmunoglobulin receptor of a first animal and which is analogous to at least a portion of the following amino acid residue segments of the rabbit polyimmunoglobulin receptor: domain I, domain II, domain III, domain IV, domain V, and amino acid residues 553 to 627 of domain VI;
  - b) at least one amino acid segment which is from the polyimmunoglobulin receptor of a second animal and which corresponds to or is analogous to the following amino acid residue segments of the rabbit polyimmunoglobulin receptor: domain I, domain II, domain III, domain IV, domain V, and amino acid residues 553 to 627 of domain VI.
  - 18. The immunoglobulin of claim 16 wherein said first animal is a mammal and said second animal is a rabbit.
  - 19. The immunoglobulin of claim 16 wherein said first animal is a human and said second animal is a rabbat.

- The immunoglobulin of claim 1 wherein said immunoglobulin derived \heavy chain contains at least a portion of an IgM or IgA heavy chain of any subtype
- The immunoglobulin of claim 1 wherein said immunoglobulin derived heavy chain is comprised of/immunoglobulin domains from two different isotopes of immunoglobulin.
- The immunoglobulin of claim 21 wherein said immunoglobulin domains are selected from the group consisting of:
  - the  $C_H1$  of a mouse IgG1 and the  $C_H2$  and  $C_H3$  of a) mouse IgA; and
  - the  $C_H1$  and  $C_{H2}$  of a mouse LGG1 and the  $C_{H2}$  and b) C<sub>H</sub>3 of mouse IgA;
  - 23. The immunoglobulin of plaim 1 wherein said antigen binding domain substantially corresponds to the Guy's 13 heavy chain variable region.
- The immunoglobu/14/n of claim 2 wherein said antigen binding domain substantially corresponds to the 20 Guy's 13 light chain variable region.
  - 25. The immunoglobulin of claim 1 wherein said protection protein has a first amino acid sequence which substantially corresponds to at least a portion of the amino acid residues 1 to 606 or 1 to 627 of the rabbit polyimmunoglobulin receptor and has a second amino acid residue sequence contiguous with said first amino acid sequence, wherein said second amino acid residue sequence does not have an amino acid residue sequence corresponding to the functional transmembrane segment of the rabbit polyimmunoglobulin receptor.
  - The /immunoglobulin of claim 25 wherein said second amino acid residue sequence has an amino acid sequence which corresponds to amino acid residues 655 to 755 of a pol/immunoglobulin receptor.
- The immunoglobulin of claim 25 wherein said second amino acid residue sequence is a portion of one or more of the following: an intracellular domain of a

polyimmunoglobulin molecule, a domain of a member of the immunoglobulin gene superfamily, an enzyme, a toxin, or a

linker.

Suction 28. A eukaryotic cell containing an immunoglobulin of claims 1-24.

29. The eukaryotic cell of claim 28 wherein said eukaryotic cell is a plant cell.

The plant cell of claim 29 wherein said plant cell is part of a plant.

31. A eukaryotic cell containing a nucleotide sequence encoding a protection protein.

32. The eukaryotic cell of claim 31 which also contains a second nucleotide sequence encoding at least one of the molecules selected from the group consisting of: an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain, an immunoglobulin derived light chain having at least a portion of an antigen binding domain, or an immunoglobulin J chain.

33. The eukaryotic cell of claim 32 wherein said second nucleotide sequence encodes an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain; and which also contains a third nucleotide sequence encoding an immunoglobulin derived light chain having at least a portion of an antigen binding domain.

34. The eukaryotic cell of claim 33 which also contains a fourth nucleotide sequence encoding an immunoglobulin J chain.

35. The emkaryotic cell of claims 31-34 wherein 30 said eukaryotic cell is a plant cell.

36. A plant cell containing a nucleotide sequence encoding a protection protein and a nucleotide sequence encoding an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain.

37. A eukaryotic cell containing a protection protein.

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- 38. The eukaryotic cell of claim 37 which also contains at least one additional molecule selected from the group consisting of: an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain, an immunoglobulin derived light chain having at least a portion of an antigen binding domain, or an immunoglobulin J chain.
- 39. The eukaryotic cell of claim 38 wherein said additional molecule is an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain; and which also contains an immunoglobulin derived light chain having at least a portion of an antigen binding domain.

40. The eukaryotic cell of claim 37 which also 15 contains an immunoglobulin J chain.

The eukaryotic cell of claims 37-40 wherein said eukaryotic cell is a plant cell.

wherein said plant cell is derived from a dicotyledonous or monocotyledonous plant.

43. The plant cell of claims 29, 35, 36 and 41 wherein said plant cell is derived from a solanaceous plant.

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- 44. The plant/cell of claims 29, 35, 36 and 41 25 wherein said plant cell is alfalfa cell.
  - 45. The plant cell of claims 29, 35, 36 and 41 wherein said plant cell is derived from a tobacco plant.
  - wherein said plant cell is part of a plant.
- 30 47. A composition comprising an immunoglobulin of claims 1-24 and plant macromolecules.
  - 48. The composition of claim 47 wherein the plant molecules are derived from a dicotyledonous, monocotyledonous, solanaceous, alfalfa or tobacco plant.
- 35 49. The composition of claim 47 wherein said plant molecules are ribulose bisphosphate carboxylase, light

harvesting complex, pigments, secondary metabolites/or chlorophyll.

- The composition of claim 47 wherein/said 50. immunoglobulin is present in a concentration of between 0.001% and 99% mass excluding water.
- The composition of claim 47 wherein said plant 51. macromolecules are present in a concentration of between 1% and 99% mass excluding water.
- A method of producing an immunoglobulin of 10 claims 1-24 comprising the steps of:
  - (a) introducing into a plant cell an expression vector containing a nucleotide sequence encoding a protection protein operably linked to a transcriptional promoter; and
- introducing into said plant cell an expression 15 vector containing a mucledtide sequence encoding an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain operably linked to a transcriptional promoter.
  - The method of claim 52 further comprising the 53. step of:
- introducing into said plant cell an expression (c) vector containing a nucle of tide sequence encoding an immunoglobulin derived light chain having at least a 25 portion of an antigen binding domain operably linked to a transcriptional promoter.
- The method of claims 52 or 53 further comprising the step of introducing into said plant cell an expression vector containing a nucleotide sequence encod-30 ing an immunoglobulin J chain operably linked to a transcriptional/promoter.
- The method of claims \$2-54 wherein said immunoglobulin derived heavy chain is immunoglobulin alpha chain and said immunoglobulin derived hight chain is an immuno-35 globulin kappa or lambda chain.

- 56. The method of claims 52-54 wherein said immunoglobulin derived heavy chain is comprised of portions of immunoglobulin alpha chain and immunoglobulin gamma chain.
- 57. The method of claims 52-54 wherein said plant cells are part of a plant.
  - 58. The method of claim 56 further comprising growing said plant.
- 59. The method of claims 56 or 57 wherein said plant is a dicotyledonous, monocotyledonous, solanaceous, leguminous, alfalfa or tobacco plant.
- 60. The methods of claims 52-59 wherein said immunoglobulin derived heavy chain is a chimeric immunoglobulin heavy chain.
- of. A method of producing a therapeutic immunoglobulin composition containing plant macromolecules, said
  method comprising the step of shearing under pressure a
  portion of a plant of claims 30 or 46 to produce a pulp
  containing a therapeutic immunoglobulin and plant macromolecules in an liquid derived from the apoplast or
  symplast of said plant and solid plant derived material.
  - 62. The method of claim 61 further comprising the step of separating said solid plant derived material from said liquid.
- 63. The method of claim 61 or 62 wherein said 25 portion of said plant is a leaf, stem, root, tuber, fruit or entire plant.
  - 64. The method of claim 61 wherein said shearing is accomplished by a mechanical device which releases liquid from the apoplast or symplast of said plant.
  - 65. The method of claim 62 wherein said separation is by centrifugation, settling, flocculation or filtration
  - 66. A method for producing an assembled immunoglobulin molecule having heavy, light and J chains and a protection protein complising the steps of:
  - a) introducing into a eukaryotic cell nucleotide sequences operably linked for expression encoding:

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i) an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain,

- ii) an immunoglobulin derived light chain having at least a portion of an antigen binding domain,
- iii) an immunoglobulin J chain, and
- iv) a protection protein; and
- b) maintaining said cell under conditions allowing 10 production and assembly of said immunoglobulin derived heavy and light chains, said immunoglobulin J chain and said protection protein into an immunoglobulin molecule.
  - of. A method for producing an assembled immunoglobulin molecule having heavy, light and J chains and a protection protein by maintaining under conditions allowing protein production and immunoglobulin assembly, a eukaryotic cell containing nucleotide sequences operably linked for expression encoding:

i) an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain,

- ii) an immunoglobulin derived light chain having at least a portion of an antigen binding domain,
- iii) an mmunog/Pobulin J chain, and

iv) a protection protein.

68. The method of claims 66-67 wherein said eukaryotic cell is a plant cell.

69. A method of making an immunoglobulin resistant 30 to environmental conditions comprising the steps of :

operably linking a nucleotide sequence encoding at least a portion of the antigen binding domain derived from an immunoglobulin heavy chain to a nucleotide sequence encoding at least one domain derived from an immunoglobulin alpha heavy chain to form a nucleotide sequence encoding a chimeric immunoglobulin heavy chain;

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b) expressing said nucleotide sequence encoding said chimeric immunoglobulin heavy chain to produce said chimeric immunoglobulin heavy chain in a eukaryotic cell which also contains at least one other molecule selected from the group consisting of: a protection protein, an immunoglobulin derived light chain having at least a portion of an antigen binding domain and an immunoglobulin J chain; and

thereby allowing the chimeric immunoglobulin heavy chain to assemble with said at least one other molecule to form said immunoglobulin resistant to said environmental conditions.

70. The method of claim 69 wherein said other 15 molecule is a protection protein and said eukaryotic cell also contains an immunoglobulin derived light chain having at least a portion of an antigen binding domain and an immunoglobulin J chain.

71. A process for producing an immunoglobulin 20 resistant to environmental conditions by maintaining under conditions allowing protein production and immunoglobulin assembly a cell containing:

a) a nucleotide sequence encoding a chimeric immunoglobulin heavy chain in which a nucleotide sequence encoding at least a portion of an antigen binding domain derived from heavy chain is operably linked to a nucleotide sequence encoding at least one domain derived from an immunoglobulin alpha heavy chain; and

b) at least one other molecule selected from the group consisting of: a protection protein, an immunoglobulin derived light chain having at least a portion of an antigen binding domain and an immunoglobulin J chain;

thereby allowing the chimeric immunoglobulin heavy chain to assemble with said at least one other molecule to form

said immunoglobulin fesistant to said environmental conditions.

- 72. The immunographulin of claims 50-52 wherein said eukaryotic cell is a prant cell.
- 73. A tetratransgenic organism comprised of cells containing four different transgenes each encoding a different polypeptide of a multipeptide molecule wherein at least one of each of said different polypeptides is associated together in said multipeptide molecule.
- 74. The transgenid organism of claim 73 wherein at least one of said four transgenes is a transgene encoding a protection protein.
  - 75. The transgenic organism of claim //3 wherein at least one of said four transgenes is a transgene encoding an immunoglobulin derived heavy chain having at least a portion of an antigen binding domain.
- 76. The transgenic organism of claim 73 wherein at least one of said four transgenes is a transgene encoding an immunoglobulin derived light chain having at least a 20 portion of an antigen binding domain.
  - 77. The transgenic organism of claim 73 wherein at least one of said four transgenes is a transgene encoding an immunoglobulin J chain.
- 78. The transgenic organism of claim 73 wherein at least one of said four transgenes is a transgene encoding a chimeric immunoglobulin heavy chain.
  - 79. The transgenic organism of claim 73 wherein said transgenic organism is a plant.
- 80. The transgenic organism of claim 73 wherein 30 said transgenic organism is a mammal.
  - 81. The immunoglobulin of claim 1 wherein said chimeric immunoglobulin heavy chain contains an immunoglobulin domain from one of the following immunoglobulin heavy chains: IgG, IgA, IgM, IgE, IgD; and also contains a protection protein-binding domain from IgA or IgM.
  - 82. The immunoglobulin of claim 81 wherein said immunoglobulin heavy chains are human, rodent, rabbit,

bovine, ovine, caprime, fowl, canine, feline or primate immunoglobulin heavy chains.

- 83. The immunoglobulin of claim 81 wherein said protection protein-binding domain is from the IgA of a 5 human, rodent, rabbit, bovine, ovine, canine, feline or primate.
  - 84. The immunoglobulin of claim 81 wherein said chimeric immunoglobulin heavy chain is comprised of immunoglobulin chains of mouse IgG1 and said protection protein-binding domain is from mouse IgA or IgM.
  - 85. The immunoglobulin of claim 81 wherein said chimeric immunoglobulin heavy chain is comprised of immunoglobulin domains of a human IgG, IgM, IgD or IgE and said protection protein-binding domain is from a human IgA

